## CENTRAL PAX CENTER

## IN THE CLAIMS

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1. (currently amended) A method comprising treating an allergic skin disease by topically administering for the first time after an allergic challenge to a subject in need thereof a therapeutically effective amount of a compound of formula (1) or a pharmaceutically acceptable salt thereof:

$$R^2$$
 $R^3$ 
 $R^4$ 
 $R^1$ 

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in which

 $\mathbf{R}^1$  is

(i)  $-C_{1-12}$ -alkyl, straight-chain or branched-chain or  $-C_2-C_{12}$  alkenyl, mono- or polyunsaturated,

optionally mono- or polysubstituted by -OH, -SH, -NH<sub>2</sub>, -NHC<sub>1-6</sub>-alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub> aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-14</sub>aryl)<sub>2</sub> -N(C<sub>1-6</sub>alkyl)(C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>, -NO<sub>2</sub>, -CN, -F, -Cl, -Br, -I, -O-C<sub>1-6</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>6</sup>, -OSO<sub>2</sub>C<sub>1-6</sub>-alkyl, -OSO<sub>2</sub>C<sub>6-14</sub>aryl, -(CS)R<sup>6</sup>, -COOH, -(CO)R<sup>6</sup>, mono-, bi or tricyclic

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saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or tricyclic saturated or mono-or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the  $C_{6-14}$ aryl groups and the carbocyclic and heterocyclic substituents for their part can optionally be mono- or polysubstituted by  $R^4$ .

(ii) a mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycle having 3-14 ring members or a mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycle having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, or a carbo- or heterocyclic saturated or mono- or polyunsaturated spirocycle having 3-10 ring members, where heterocyclic systems contain 1-6 heteroatoms, which are preferably N, O and S, optionally mono- or polysubstituted by -OH, -SH, -NH2, NHC<sub>1-6</sub> alkyl, -N (C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-6</sub>alkyl) (C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>-NO<sub>2</sub>, -CN, -F, -Cl, -Br, -I, -O-C<sub>1-6</sub> alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>6</sup>, OSO<sub>2</sub>C<sub>1-6</sub>alkyl, -OSO<sub>2</sub>C <sub>6-14</sub> aryl, -(CS)R<sup>6</sup>, -COOH, -(CO)R<sup>6</sup> mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the C<sub>6-14</sub>aryl groups and the carbocyclic and heterocyclic substituents can optionally be mono- or polysubstituted by R<sup>4</sup>.

R<sup>5</sup> is

a mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycle having 3-14 ring members or a mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycle having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, or a carbo-or heterocyclic saturated or mono-or polyunsaturated spirocycle having 3-10 ring members, where heterocyclic systems contain 1-6 heteroatoms, which preferably N, O and S, optionally mono- or

polysubstituted by -OH, -SH, -NH<sub>2</sub> -NHC<sub>1-6</sub> alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-6</sub>alkyl)(C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>, -NO<sub>2</sub> -CN, -F, -Cl, -Br, -I, -O-C<sub>-1-5</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>6</sup>, -OSO<sub>2</sub>C<sub>1-6</sub>alkyl, OSO<sub>2</sub>C<sub>6-14</sub>aryl, -(CS)R<sup>6</sup>, -COOH, -(CO)R<sup>6</sup>, mono-, bi or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the C<sub>6-14</sub>aryl groups and the carbocyclic and heterocyclic substituents can optionally be mono- or polysubstituted by R<sup>4</sup> with the proviso that R<sup>5</sup> contains at least one substituent selected from -F, -Cl, -Br, -I,

 $R^2$ ,  $R^3$  are hydrogen or -OH, where at least one of the two substituents must be -OH;  $R^4$  is

-H, -OH, -SH, -NH<sub>2</sub> -NHC<sub>1-6</sub>-alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-6</sub> alkyl) (C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>, -NO<sub>2</sub>, -CN, -COOH, - (CO)R<sup>6</sup>, -(CS)R<sup>6</sup>, -F, --Cl, -Br, -I, -O-C<sub>1-6</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>2</sub>R<sup>6</sup>, -C<sub>1</sub>-C<sub>6</sub>-alkyl, wherein each aryl or alkyl may be mono- or polysubstituted by -OH, -F, -Cl, -Br, -I;

R<sup>6</sup> is

-H, -NH<sub>2</sub>, -NHC<sub>1-6</sub>-alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, - N(C<sub>1-6</sub>alkyl) (C<sub>6-14</sub>aryl), -O-C<sub>1-6</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl,

-C1-12-alkyl, straight-chain or branched-chain,

-C2-12-alkenyl, mono- or polyunsaturated, straight-chain or branched-chain,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S;

A is either a bond, or

$$-(CH_2)_m -(CH=CH)_n -(CH_2)_p - (CHOZ)_m - (C=O) - (C=S) - (C=N-Z) - O- (C=N-Z) - (C=N-Z) - O- (C=N-Z) - (C=N-Z) -$$

wherein m, p=0-3 and n=0-2 and

Z is

-H, or

-C<sub>1-12</sub>-alkyl, straight-chain or branched-chain,

-C2-12-alkenyl, mono- or polyunsaturated, straight-chain or branched-chain,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S;

B is either carbon or sulfur, or -(S=O)-;

D is O, S, CH<sub>2</sub> or N-Z,

where, if B is carbon, D is O, S or CH2;

E is a bond, or

-(CH<sub>2</sub>)<sub>m</sub>-, -0-, -S-, -(N-Z)-, wherein m and Z have the meaning already described above.

2. (previously presented) The method of claim 1 wherein R<sup>5</sup> is selected from monocyclic saturated or mono- or polyunsaturated carbocycles and heterocycles having at least one halogen substituent.

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- 3. (previously presented) The method of claim 2 wherein R<sup>5</sup> is selected from monocyclic aromatic carbocycles and heterocycles having at least one halogen substituent.
- 4. (previously presented) The method of claim 3 wherein R<sup>5</sup> is a pyridine ring having at least one halogen substituent.
  - 5. (canceled)
- 6. (previously presented) The method of claim 1 wherein  $R^1$  is selected from  $C_1$ - $C_{12}$  alkyl, which is optionally substituted.
  - 7. (canceled)
  - 8. (previously presented) The method of claim I wherein  $R^2$  is OH and  $R^3$  is H.
- 9. (previously presented) The method of claim 1 wherein A is selected from -(C=O)-and -(CHOH)-.
  - 10. (previously presented) The method of claim 1 wherein B is C.
  - 11. (previously presented) The method of claim 1 wherein D is O.
  - 12. (previously presented) The method of claim 1 wherein E is -(N--H)-.
- 13. (previously presented) The method of claim 1 wherein compound (I) is (N-3,5-dichloro-4-pyridinyl)-2-[1-(4-fluorobenzyl)-5-hydroxy-1 H-indol-3-yl]-2-oxoacetamide).
  - 14. (canceled)
- 15. (previously presented) The method of claim 1 wherein the disease is allergic dermatitis.
  - 16. (canceled)
- 17. (previously presented) The method of claim 1, wherein the compound is administered to a skin area which is afflicted with the disease.

- 18. (previously presented) The method of claim 17 wherein the compound is administered up to 48 h after the allergic challenge.
  - 19. (canceled)
- 20. (currently amended) The method of claim 1, claim 1 wherein a further pharmaceutical agent is administered, wherein said further pharmaceutical agent stimulates cAMP production and is selected from the group consisting of a sympathomimetic amine, a xanthine derivative, a corticosteroid and an adrenal stimulant.
- 21. (previously presented) The method of claim 20 wherein the further pharmaceutical agent is a corticosteroid.
- 22. (previously presented) The method of claim 20, wherein the allergic disease is allergic dermatitis.
- 23. (previously presented) The method of claim 1, further comprising administering a further pharmaceutical agent.